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RAW SEQUENCE LISTING

PATENT APPLICATION: US/09/903,325A

DATE: 03/13/2002 TIME: 11:17:12

Input Set : A:\510015-257.TXT

Output Set: N:\CRF3\03132002\I903325A.raw

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4 <110> APPLICANT: De Robertis, Edward M.
              Bouwmeester, Tewis
     8 <120> TITLE OF INVENTION: Endoderm, Cardiac and Neural Inducing
             Factors
     11 <130> FILE REFERENCE: 510015-257
     13 <140> CURRENT APPLICATION NUMBER: US 09/903,325A
C--> 14 <141> CURRENT FILING DATE: 2001-11-07
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    17 <151> PRIOR FILING DATE: 1996-06-20
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    26 <213> ORGANISM: Xenopus
    28 <400> SEQUENCE: 1
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        Ser Leu Asn Ser Arg Gly Tyr Phe Arg Lys Glu Arg Gly Ala Arg Arg
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    34
        Ser Lys Ile Leu Leu Val Asn Thr Lys Gly Leu Asp Glu Pro His Ile
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    36
                                55
       Gly His Gly Asp Phe Gly Leu Val Ala Glu Leu Phe Asp Ser Thr Arg
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    38
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                                                 75
       Thr His Thr Asn Arg Lys Glu Pro Asp Met Asn Lys Val Lys Leu Phe
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       Ser Thr Val Ala His Gly Asn Lys Ser Ala Arg Arg Lys Ala Tyr Asn
   42
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       Gly Ser Arg Arg Asn Ile Phe Ser Arg Arg Ser Phe Asp Lys Arg Asn
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       Thr Glu Val Thr Glu Lys Pro Gly Ala Lys Met Phe Trp Asn Asn Phe
   45
   46
                                135
       Leu Val Lys Met Asn Gly Ala Pro Gln Asn Thr Ser His Gly Ser Lys
   47
   48
                           150
                                                155
       Ala Gln Glu Ile Met Lys Glu Ala Cys Lys Thr Leu Pro Phe Thr Gln
   49
   50
                       165
                                            170
       Asn Ile Val His Glu Asn Cys Asp Arg Met Val Ile Gln Asn Asn Leu
   51
   52
                                        185
   53
       Cys Phe Gly Lys Cys Ile Ser Leu His Val Pro Asn Gln Gln Asp Arg
   54
               195
                                   200
       Arg Asn Thr Cys Ser His Cys Leu Pro Ser Lys Phe Thr Leu Asn His
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56
          210
                             215
                                                 220
     Leu Thr Leu Asn Cys Thr Gly Ser Lys Asn Val Val Lys Val Val Met
  57
  58
                         230
                                             235
     Met Val Glu Glu Cys Thr Cys Glu Ala His Lys Ser Asn Phe His Gln
  59
  60
                                         250
     Thr Ala Gln Phe Asn Met Asp Thr Ser Thr Thr Leu His His
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  62
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  67 <213> ORGANISM: Xenopus
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     atgtactcag gatctgtatt atcgtctgcc ttgtgaatga tggagcagga aaacactcag
                                                                            60
     120
 73
    gaggagcacg taggagcaag attctgctgg tgaatactaa aggtcttgat gaaccccaca
                                                                           180
    ttgggcatgg tgattttcgc ttagtagctg aactatttga ttccaccaga acacatacaa
                                                                           240
 74
     acagaaaaga gccagacatg aacaaagtca agcttttctc aacagttgcc catggaaaca
                                                                           300
 75
    aaagtgcaag aagaaaagct tacaatggtt ctagaaggaa tatttttcct cgccgttctt
                                                                           360
 77
     ttgataaaag aaatacagag gttactgaaa agcctggtgc caagatgttc tggaacaatt
                                                                           420
 78
     ttttggttaa aatgaatgga gccccacaga atacaagcca tggcagtaaa gcacaggaaa
                                                                          480
     taatgaaaga agcttgcaaa accttgtttt tcactcagaa tattgtacat gaaaactgtg
                                                                          540
 79
    acaggatggt gatacagaac aatctgtgct ttggtaaatg catctctctc catgttccaa
                                                                          600
 80
    atcagcaaga tcgacgaaat acttgttccc attgcttgcc gtccaaattt accctgaacc
                                                                          660
 81
    acctgacgct gaattgtact ggatctaaga atgtagtaaa ggttgtcatg atggtagagg
                                                                          720
 82
    aatgcacgtg tgaagctcat aagagcaact tccaccaaac tgcacagttt aacatggata
                                                                          780
    catctactac cctgcaccat taaaggactg ccatacagta tggaaatgcc cttttgttgg
                                                                          840
 84
    aatatttgtt acatactatg catctaaagc attatgttgc cttctatttc atataaccac
                                                                          900
 85
    atggaataag gattgtatga attataatta acaaatggca ttttgtgtaa catgcaagat
                                                                          960
    ctctgttcca tcagttgcaa gataaaaggc aatatttgtt tgactttttt tctacaaaat
                                                                         1020
 87
    gaatacccaa atatatgata agataatggg gtcaaaactg ttaaggggta atgtaataat
                                                                         1080
88
    agggactaag tttgcccagg agcagtgacc cataacaacc aatcagcagg tatgatttac
                                                                         1140
89
    tggtcacctg tttaaaagca aacatcttat tggttgctat gggttactgc ttctgggcaa
                                                                         1200
    aatgtgtgcc tcataggggg gttagtgtgt tgtgtactga ataaattgta tttatttcat
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92
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95 <211> LENGTH: 318
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97 <213> ORGANISM: Xenopus frazzled
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                     5
                                                            15
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102
103
    Pro Val Arg Ile Pro Met Cys Lys Ser Met Pro Trp Asn Met Thr Lys
104
105
                                40
    Met Pro Asn His Leu His His Ser Thr Gln Ala Asn Ala Ile Leu Ala
106
107
                            55
    Ile Glu Gln Phe Glu Gly Leu Leu Thr Thr Glu Cys Ser Gln Asp Leu
108
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T. T	
109 65 70 75	
110 Leu Phe Phe Leu Cys Ala Met Tyr Ala Pro 75	
110 Leu Phe Phe Leu Cys Ala Met Tyr Ala Pro Ile Cys Thr Ile Asp Phe	
112 Gln His Glu Pro Ile Lys Pro Cya Lys 90	
112 Gln His Glu Pro Ile Lys Pro Cys Lys Ser Val Cys Glu Arg Ala Arg	
114 Ala Gly Cys Glu Pro Ile Iou Ile I	
114 Ala Gly Cys Glu Pro Ile Leu Ile Lys Tyr Arg His Thr Trp Pro Glu	
116 Ser Leu Ala Cys Glu Clu Lau D 125	
116 Ser Leu Ala Cys Glu Glu Leu Pro Val Tyr Asp Arg Gly Val Cys Ile	
118 Ser Pro Glu Ala Ilo Val 71 140	
118 Ser Pro Glu Ala Ile Val Thr Val Glu Gln Gly Thr Asp Ser Met Pro 129 145 150 150	
120 Asp Phe Ser Mot Asp Care 155	
120 Asp Phe Ser Met Asp Ser Asn Asn Gly Asn Cys Gly Ser Gly Arg Glu 165 170	
122 His Cys Lys Cys Lys Pro W 1 7 170 175	
122 His Cys Lys Cys Lys Pro Met Lys Ala Thr Gln Lys Thr Tyr Leu Lys	
180 185 190 185 190	
124 Asn Asn Tyr Asn Tyr Val Ile Arg Ala Lys Val Lys Glu Val Lys Val 125 200 200	
126 Lys Cys His Asp Ala Why 13	
126 Lys Cys His Asp Ala Thr Ala Ile Val Glu Val Lys Glu Ile Leu Lys 205 217 210 215	
128 Ser Ser Leu Val	
and the ser ber ber val Ash The Pro Lyg Ash mi	
129 225 230 235 230 235	
240 Del Gly CVS Lell Cvg Dro Clasta and 240	
131 245 250 250 255 250 255	
132 Het Gry Tyr Glu Asp Lys Glu Arg mb- 2	
133 260 265 265	
134 Ser Leu Ala Glu Lys Tro Arg Aco Are 1	
135 275 280 Leu Ala Lys Lys Val Lys Arg	
136 Trp Asp Gln Lys Leu Arg Arg Bro Arg 7	
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142 <211> LENGTH: 1875	
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149 Ctaattctqc actititaaat tatotaatta sagatuggat tydaggactt ggatttttat	120
150 gataaactta actoottaat tttaaattaaa	180
151 ttgcttttac atgtgcccag attttag Coddadata Laaggtgggg tgagttgtag	240
152 acacatacag gttgggcaga ataggata	300
153 tactggccat acctggactg gggettatal occadaday gaaagtggac tcattactgc	360
154 agectgtgcg gatccccatg tages to the state of the taget act act act act act act act act act ac	420
155 atctccacca cagcactcaa goodata discouraged catgaccaag atgcccaacc	480
156 tgaccactga atgtagccag gacattti	540
156 tgaccactga atgtagccag gaccttttgt tetttetgtg tgecatgtat gaaggtttge 157 gtaccatega tttecagcat gaaccaatta agecttgaa gtacat	600
157 gtaccatcga tttccagcat gaaccaatta agccttgcaa gtccgtgtgc gaaagggcca	
158 gggccggctg tgagcccatt ctcataaagt accggcacac ttggccagag agcctggcat 159 gtgaagagct gcccgtatat gacagaggag tctgcatata gacagag agcctggcat	660
159 gtgaagaget geoogtatat gacagaggag tetgeatete eccagagget ategteacag	720
	780

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160	\ +~.																
160		Jaac	aagg	aac	agat	tca	atgc	caga	.ct t	ctcc	atgg	a tt	caaa	caat	gga	aattgcg	840
161	. yac	iyeg	gcag	gga	gcac	tgt	aaat	qcaa	gc c	catq	aaga	с аа	ccca	aaaa	aco	tatotos	900
162	aya	iala	atta	ggagcactgt aaatgcaagc ccatgaaggc aacccaaaag acgtatctca caattatgta atcagagcaa aagtgaaaga ggtgaaagtg aaatgccacg										960			
163	ace	Juaa	cage	deligiggaa qtaaaqqaqa ttotoaagto ttoootagto aacattoota										1020			
164	aag	jaca	cayı	gacactgrac accaactcag gctgcttgtg cccccagett gttgccaatg											1080		
165	ayy	jaat	acai	. dattatgggc tatgaagaca aagagcgtac caggcttota otagtggaag													
166	gat	cct	tggc	cga	aaaa	tgg	agag	atco	te t	tact	ааса	a am	tcaa	acac	taa	gcggaag	1140
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168	att	cca	gaca	agc	acat	aαt	taσa.	ctaa	ca a	aaam	atat	c aa	~~~~	caac	aaa	gactttg	1260
169	aaa	cta	aσat.	t.t.a	catt	att	agaa	Tago	29 9	2220	222+	a Ly	yaaa	CLCL	atg	gactttg ttatatt	1320
170	cta	t.t.a	ttta	cta	caad	aan	ctaa	+++>	au a	~~++	aaat	L go	acta	cage	acg	ttatatt	1380
171	tta	taad	rtat	att	taas	aat .	atta	2222	96 6	gall	grag	t tc	rcct.	ttcc	ttc	tttttt	1440
172		taa	rtaa	ato	tata	og c	gtto	cay	ge a	actg		a tt	caac	ttcc	agt	gacagag	1500
173		tati	ta	+20	LU LU:	age	CLda	agaa	ge to	caat	tcat	t tc	tgat	caac	taa	tggtgac	1560
174	022	+4+	-cya	Lac	LLGG	yya i	aagto	gaac	ca ai	ttgc	aatg	agag	aaa	agttgac	1620		
175		+++	-gci		ectg.	cag a	atgaa	acaa	gt ga	agaga	atcad	c at	ttaaa	atga	tga	tcacttt	1680
	CCa	LLLC	iala	CLL	ccago	cag 1	tttta	agtta	acrat	cgaca	atata	a aaa	atac:	acct	2221	totaaat	1740
176	all	LLai	-cat	aaaı	.gaaq	gag (ctqqt	ittad	aa ct	cotai	.aat.o	r act	atta	annr	agg	taaataa	1800
177	CLa		-y cc	aati	cctgi	ttt 1	taaaa	atto	gc ct	aaat	aaat	t att	aagt	cct	aaat	taaaaaa	1860
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187	1				5	-	,			10	. neu	r neu	. Leu	с Сту		и мет	
188	Val	Leu	Gln	Thr	Asr	Cvs	Glu	Tla	Δla	Gla	Ттт	. m	т1.	3	15	Glu	
189				20		012	Olu	. 110	25	GIII	тут	т Ат	тте		Glu	ı Glu	
190	Glu	Pro	Pro		Thr	Val	Tlo	λ 1 -		т о	a			30		. Phe	
191			35	0-1		val		40	vai	Leu	ser	GIN		Ser	Ile	Phe	
192	Asn	Thr		Aen	Tlo	Dro	. ala			5 1	_	_	45				
193	11011	50	T 11T	nsp	тте	PIO	Ald	Thr	Asn	Phe	Arg		Met	Lys	Gln	Phe	
194	λen		Com	т	T1 -	a 1	55	_				60					
195	65	USII	ser	ьeu	тте	GIY	val	Arg	GLu	Ser	Asp	Gly	Gln	Leu	Ser	Ile	
196		a 1		- 1	_	70					75					80	
	мес	GIU	Arg	тте	Asp	Arg	Glu	Gln	Ile	Cys	Arg	Gln	Ser	Leu	His	Cys	
197					85					90					95	_	
198	Asn	Leu	Ala	Leu	Asp	Val	Val	Ser	Phe	Ser	Lys	Gly	His	Phe	Lys	Leu	
199				TOO					105					110			
200	Leu	Asn	Val	Lys	Val	Glu	Val	Arg	Asp	Ile	Asn	Asp	His	Ser	Pro	His	
201			TTJ					120					125				
202	Phe	Pro	Ser	Glu	Ile	Met	His	Val	Glu	Val	Ser	Glu	Ser	Ser	Ser	Wa 1	
203		T20					135					140					
204	Gly	Thr	Arg	Ile	Pro	Leu		IJe	Ala	Tle	Acn	Glu	Δαη	Wa 1	C1	Com	
205	145		-			150					155	oru.	ռոր	val	GTÀ		
206	Asn :	Ser	Ile	Gln	Asn	Phe	Gln	Tlο	Ser	Δαη	T 2 2	Co~	114 -	nk -	α.	160	
207		_			165		0111	116	PEI	170	ASII	ser	HIS	rne		īīe	
208	Asp '	Va l	Len	Thr		Δla	λαη	C1.,	17 - 1	170	m	n 7	_	_	175		
209	Asp '		cu	180	y	Ala	vah	GTĀ	10E	гÀ2	туr	Ата	Asp		Val	Leu	
210	Met i	۵ra	Glu		λας	7 m~	C1	т1 -	185	_	_,	_	_	190			
•	Met A	9	JIU	⊒-cu	ush	лту	GIU	тте	GIN	Pro	Inr	Tyr	ITe	Met	Glu	Leu	

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21				19						. 2	000											
21	2 L	eu .	Ala	Me	t A	sp G	lv (G1 v	. Wa	ם די	200		,	-	_		_ :	205	5			
21:	3		210			-r •	-1	o i y	21	2 1.T. E	10	56	er l	Leu	Se	r G.	Ly '	Thr	A]	a v	al	Val
214	4 A.	sn	Ile	Ar	αV	al I.	211 2	N con	Z.I Dh				_	_		2:	20					
215	5 2	25			J ''	~ ± 10	cu i	230	PII	le A	sn	As	p A	Asn	Se	r Pi	ro I	Val	. Ph	ie G	lu	Arg
216																						
217	7			11	C A.	La Va	11 <i>E</i>	ısp	ье	u v	aΙ	G1	u A	lsp	Al	a Pı	o I	Leu	Gl	у Т	yr	Leu
218																						
219)	-u 1	Jeu	GI	T THE	eu H: 50	LS A	та	Th	r A	sp	As	p A	sp	Glı	u Gl	y V	al	As	n G	lv	Glu
220																						
221	, 11		ат	7 J J	- c. G1	y Pł	ie S	er	Th:	r L	eu	Al:	a S	er	Gli	a Gl	u V	al	Ar	a G1	ln	I.eu
222																						
223	P1.	ie ī	iys	116	As	n Se	er A	rg	Th:	r G	lу	Se	r V	al	Thi	Le	u G	lu	G1	v G1	n	Val
223																						
	AS	ЪЪ	ne	Glu	ιTh	r Ly	s G	ln	Thi	r Ty	/r	Glu	ı P	he	Glu	ı Va] G	l n	Δ1:	a C1	n	7.00
225																						
226	гe	u G	TÀ	Pro	As	n Pr 32	o L	eu	Thi	: Al	a	Thr	. C	vs	Lvs	. Va	יתי ו	hr	170	1 114	_	320
227																						
228	Le	u A	sp	Val	As	n As	рΑ	sn	Thr	Pr	0	Ala	ı T	l e	Thr	т1.	о mi	h m	D	33	5	_,
229																						
230	Th	r V	al	Asn	Ala	a Gl	y Va	al	Ala	Tv	r	Tle	D:	ro	c1n	mh.	. a.	1 _	350	, _		_
231																						
232	Ası	n Pi	he	Ile	Ala	a Le	ı I.	le	Ser	· Th	r	Thr	. λ.	'n	7 ~~	37.		55				
233																						
234	Gly	γ G.	ln '	Val	Arc	ј Су:	s Tł	ır	Len	Ψъ	r (~ 1	11.4	_	~ 1	380)					
235	385	5			-	,	39	20	u	т у.	Ι (вту	п	.S (GIU	His	Ph	ıe	Lys	Lei	1 (Sln
236	Glr	ı A]	la :	Tyr	Glu	Asp	Se	י יינג	Tur	Mo	. -	т1 а	17-	1 .	395						4	100
237																						
238	Arc	r Gl	u A	Asn	Tle	Ala	י ומ	a '	Ptr~	Co.	. T		41	.0						415	5	
239					420			.u .	т Ут	se.	. 1	-eu	Tn	r \	/al	Val	Al	a (Glu	Asp	L	eu
240																						
241	_		4	135	DCI	Leu	. цу	5 .	r III T.	гуз	5 L	₁ys	Ту	r 1	'yr	Thr	۷a	1 1	Lys	Val	S	er
242																						
243	•	45	0		шъ	Asn	ΑI	a i	LO	vaı	- P	he	Se.	r I	ıуs	Pro	G1	n I	ľyr	Glu	A	la
244																						
245	465			ıcu	Giu	Asn	AS	n A	та	Pro	G	ly	Se:	r T	yr	Ile	Th	ŕ 1	hr	Val	I	le
246																						
247		***	9 5	ъp	ser	Asp	se:	r A	sp	Gln	A	sn	Gly	γL	уs	Val	Ası	n n	'yr	Ara	Le	913
248																						
249	141	A.S	PA	ıa.	LYS	Val	Met	t G	ŢУ	Gln	S	er	Leu	1 T	hr	Thr	Phe	e V	al	Ser	T.e	211
250																						
251	лэр	ATO	1 A.	sp :	ser	Gly	Va]	L	eu	Arg	A.	la	Va]	. A	rg :	Ser	Let	1 A	sp	Tvr	G1	111
252	T ***	т	Э. - -	12						520					_		525	;		-1-	U,	·u
253	пЛЯ	rec	י די	ys (31n	Leu	Asp	P.	he (Glu	I.	le (Glu	Α.	la Z	Ala	Asp) Δ	s n	G1 v	т1	0
254	Pro 545	GIR	ı Le	eu 5	Ser	Thr	Arg	Va	al (Gln	Le	eu <i>i</i>	Asn	Lε	eu 7	۱ra	T۱۵	17:	a 7	7 02	۵1	n
256	Asn	Asp	As	sn C	ys	Pro	Val	I	le :	Thr	As	sn I	?ro	J.e	- 211 T	,e11	Δen	λ.	n í	71	26	U
258	Gly	Glu	Va	ıl L	eu	Leu	Pro	IJ	le S	Ser	A .1	a F	ro	G1	n A	gn	ጥ፣፣•~	т.		7-1		
259				5	80						58	5		J		211	т Л Г.			/al	Рħ	e
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L:14 M:271 C: Current Filing Date differs, Replaced Current Filing Date